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**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicants : Rina Aharoni et al. Examiner: A. De Cloux

U.S. Serial No.: 09/768,872 Group Art Unit: 1644

Filed : January 23, 2001

For : TREATMENT OF AUTOIMMUNE CONDITIONS WITH COPOLYMER 1 RELATED COPOLYMERS AND PEPTIDES.

1185 Avenue of the Americas
New York, New York 10036
June 26, 2002

Assistant Commissioner for Patents
Washington, D.C. 20231

SIR:

**INFORMATION DISCLOSURE STATEMENT**
**PURSUANT TO 37 C.F.R. §1.97(b)(3)**

In accordance with their duty of disclosure under 37 C.F.R. §1.56, applicants direct the Examiner's attention to the following Reference Items 1-159 **(Exhibits 1-149)** which are listed again on the accompanying Form PTO-1449 **(Exhibit A)**. Applicants request that the Examiner review the references and make them of record in the subject application.

This Information Disclosure Statement is being submitted before the issuance of a first Office Action on the merits in connection with the subject application. Accordingly, no fee is required and this Information Disclosure Statement shall be considered pursuant to 37 C.F.R. §1.97(b)(3).

For the convenience of the Examiner, applicants point out that Reference Item 108 was cited in the October 29, 1999 International Search Report in the corresponding PCT International Application, and a copy of the Report is enclosed as **Exhibit B**.

Applicants also point out that several of the listed references are

Applicants also point out that several of the listed references are counterparts of each other and are cumulative. Therefore, in accordance with 37 C.F.R. § 1.98(c), a counterpart of a reference is identified after the cite to the reference, but a copy of only one of the counterparts is being provided. Applicants will provide the Examiner with copies of any reference upon request.

1. U.S. Patent No. 3,849,550, issued November 19, 1974 (Teitelbaum, et al.) **(Exhibit 1)**;

2. U.S. Patent No. 4,339,431, issued July 13, 1982 (Gaffar) **(Exhibit 2)**;

3. U.S. Patent No. 5,204,099, issued April 20, 1993 (Barbier, et al.) **(Exhibit 3)**;

4. U.S. Patent No. 5,591,629, issued January 7, 1997 (Rodriguez et al.) **(Exhibit 4)**;

5. U.S. Patent No. 5,627,206, issued May 6, 1997 (Hupe, et al.) **(Exhibit 5)**;

6. U.S. Patent No. 5,668,117, issued September 16, 1997 (Shapiro et al.) **(Exhibit 6)**;

7. U.S. Patent No. 5,719,296, issued February 17, 1998 (Acton, III, et al.) **(Exhibit 7)**;

8. U.S. Patent No. 5,800,808, issued September 1, 1998 (Konfino, et al.) **(Exhibit 8)**;

9. U.S. Patent No. 5,858,964, issued January 12, 1999 (Aharoni, et al.) **(Exhibit 9)**;

10. U.S. Patent No. 5,981,589, issued November 9, 1999 (Konfino, et al.) **(Exhibit 10)**;

11. U.S. Patent No. 5,958,972, issued September 28, 1999 (Hupe, et al.) **(Exhibit 11)**;

12. U.S. Patent No. 6,048,898, issued April 11, 2000 (Konfino, et al.) **(Exhibit 12)**;

13. U.S. Patent No. 6,054,430, issued April 25, 2000 (Konfino, et al.) **(Exhibit 13)**;

14. U.S. Patent No. 6,214,791, issued April 10, 2001 (Arnon, et al.) **(Exhibit 14)**;

15. U.S. Patent No. 6,342,476, issued January 29, 2002 (Konfino, et al.) **(Exhibit 15)**;

16. U.S. Patent Publication No. US-2001-0055568-A1, published December 27, 2001 (Gilbert et al.) **(Exhibit 16)**;

17. U.S. Serial No. 09/359,099, filed July 22, 1999 (Strominger et al.) **(Exhibit 17)**;

18. U.S. Serial No. 09/405,743, filed September 24, 1999 (Gad et al.) **(Exhibit 18)**;

19. U.S. Serial No. 09/816,989, filed March 23, 2001 (Gad et al.). Applicants point out that this reference is a counterpart of U.S. Serial No. 09/405,743(Exhibit 18);

20. U.S. Serial No. 09/875,429, filed June 5, 2001 (Yong and Chabot) **(Exhibit 19)**;

21. U.S. Serial No. 09/885,227, filed June 20, 2001 (Rodriguez and Ure) **(Exhibit 20)**;

22. PCT International Application No. PCT/US88/02139 (WO 88/10120), published December 29, 1988 (Weiner et al.) **(Exhibit 21)**;

23. PCT International Application No. PCT/US95/06551 (WO 95/31990), published November 30, 1995 (Konfino et al.). Applicants point out that this reference is a counterpart of U.S. Patents Nos. 5,800,808 (Exhibit 8) and 6,342,476 (Exhibit 15);

24. PCT International Application No. PCT/EP95/02125 (WO/95/33475), published December 14, 1995 (Kott et al.) **(Exhibit 22)**;

25. PCT International Application No. PCT/US98/00375 (WO 98/30227), published July 16, 1998 (Arnon et al.). Applicants point out that this reference is a counterpart of US Patent No. 6,214,791 (Exhibit 14);

26. PCT International Application No. PCT/US99/16617 (WO 00/05249)

published February 3, 2000 (Strominger et al). Applicants point out that this reference is a counterpart of U.S. Serial No. 09/359,099 (Exhibit 17);

27. PCT International Application No. PCT/US99/16747 (WO 00/05250) published February 3, 2000 (Aharoni et al.). Applicants point out that this reference is a counterpart of the subject application;

28. PCT International Application No. PCT/US99/22402 (WO 00/18794) published April 6, 2000 (Gad, et al.). Applicants point out that this reference is a counterpart of U.S. Serial No. 09/405,743 (Exhibit 18) and U.S. Serial No. 09/816,989;

29. PCT International Application No. PCT/US99/22836 (WO 00/20010) published April 13, 2000 (Flechter, et al.) **(Exhibit 23)**;

30. PCT International Application No. PCT/US99/27107 (WO 00/27417) published May 18, 2000 (Aharoni et al.)**(Exhibit 24)**;

31. PCT International Application No. PCT/US01/05198 (WO 01/60392) published August 23, 2001 (Gilbert et al.) Applicants point out that this reference is a counterpart of U.S. Patent Publication No. US-2001-0055568-A1 (Exhibit 16);

32. PCT International Application No. PCT/US01/18248 (WO 01/93828) published December 13, 2001 (Yong and Chabot). Applicants point out that this reference is a counterpart of U.S. Serial No. 09/875,429 (Exhibit 19);

33. PCT International Application No. PCT/US01/19649 (WO 01/97846) published December 27, 2001 (Rodriguez and Ure). Applicants point out that this reference is a counterpart of U.S. Serial

No. 09/885,227 (Exhibit 20);

34. European Patent Application No. 0 383 620 A2, published August 22, 1990 (Cook) **(Exhibit 25)**;

35. European Patent No. 0 359 783 B1, published November 29, 1995 (Werner, et al.);

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99. Meiner, "COP-1 Multicenter Clinical Trial in Exacerbating-remitting Multiple-Sclerosis: One Year Follow-up", J. Neurol., 1991(Suppl. 1) (Abstract) **(Exhibit 89)**;

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125. Fridkis-Hareli, et al., "Copolymer 1 Displaces MBP, PLP and MOG, but Can Not be Displaced by these Antigens from the MHC Class II Binding Site", Department of Chemical Immunology, The Weizmann Institute of Science, 1994 **(Exhibit 115)**;

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148. Reilly, Jr., W.J., "Pharmaceutical Necessities" in The Science and Practice of Pharmacy, Remington 1995, 2, 1380-1416 **(Exhibit 138)**;

149. Schlegel, et al., "Inhibition of Allorecognition and Prevention of Graft-vs-host Disease (GVHD) by GLAT, a Synthetic Polymer with Promiscuous Binding to Murine and Human MHC Class II Molecules", in Am. Soc. Hematology, 37th Annual Meeting, Seattle, WA (USA), December 1-5, 1995, 224a (Abstract) **(Exhibit 139)**;

150. Ben-Nun, et al., "The Autoimmune Reactivity to Myelin Oligodendrocyte Glycoprotein (MOG)in Multiple Sclerosis is Potentially Pathogenic: Effect of Copolymer 1 on MOG-induced Disease", J. Neurol., 1996, 243(Suppl. 1), S14-S22 **(Exhibit 140)**;

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152. Sykes, "Immunobiology of Transplantation", Faseb J., 1996, 10, 721-730 **(Exhibit 142)**;

153. Teitelbaum, et al., "Copolymer 1 Inhibits Chronic Relapsing

Experimental Allergic Encephalomyelitis Induced by Proteolipid Protein (PLP) Peptides in Mice and Interferes with PLP-specific T Cell Responses", J. Neuroimmunol., 1996, 64, 209-217 **(Exhibit 143)**;

154. Aharoni, et al., "Studies on the Mechanism and Specificity of the Effect of the Synthetic Random Copolymer GLAT on Graft-versus-Host Disease", Immunol. Letters, 1997, 58, 79-87 **(Exhibit 144)**;

155. Puri et al., "Modulation of the Immune Response in Multiple Sclerosis", J. Immunol., 1997, 158, 2471-2476 **(Exhibit 145)**;

156. Tarcic et al., "Copolymer 1 (Copaxone) from an Idea to a Drug for Treatment of Multiple Sclerosis" Database HCAPLUS on STN, Israel: AN 1997:333270. Kim, Handasa Kim, 1997, 281(14), 16-18 (Abstract) **(Exhibit 146)**;

157. Teitelbaum, et al., "Copolymer 1 from the Laboratory to FDA", Israel J. Med. Sci., 1997, 33, 280-284 **(Exhibit 147)**;

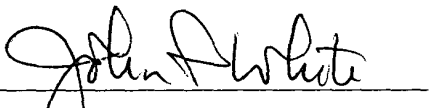
158. Fridkis-Hareli, et al., "Promiscuous Binding of Synthetic Copolymer 1 to Purified HLA-DR Molecules", J. Immunol., 1998, 160, 4386-4397 **(Exhibit 148)**; and

159. Durelli, "Immunotherapeutics of Multiple Sclerosis", Instituto di Clinica delle Malattie del Sistema Nervoso Universita di Torino, 467-475 **(Exhibit 149)**.

If a telephone conference would be of assistance in advancing the prosecution of the subject application, applicants' undersigned attorney invites the Examiner to telephone at the number provided below.

No fee is deemed necessary in connection with the filing of this Information Disclosure Statement. However, if any fee is required, authorization is hereby give to charge the amount of such fee to Deposit Account No. 03-3125.

Respectfully submitted,



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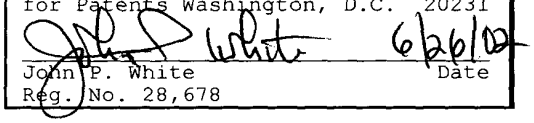
I hereby certify that this correspondence is being deposited this date with the U.S. Postal Service with sufficient postage as first class mail in an envelope addressed to: Assistant Commissioner for Patents Washington, D.C. 20231



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